

Department of Health and Human Services

Guidance on the suggested use of medical cannabis

Crohn's and Ulcerative Colitis

About this document: The following information on the use of medical cannabis serves as a suggested use guide for those participating in the Utah Medical Cannabis Program. The intended audience for this document includes qualified medical providers, pharmacy medical providers, patients intending to use medical cannabis, and caregivers of patients intending to use medical cannabis.

This document details the guidance on the use of medical cannabis for multiple sclerosis. This document does not include general instructions on the use of medical cannabis, contraindications, warnings, precautions and adverse reactions to using cannabis and drug-to-drug interactions which could be found in the extended guidance document titled *Guidance on the Suggested Use of Medical Cannabis*. The extended guidance document can be found on the Department of Health and Human Services Center for Medical Cannabis website (www.medicalcannabis.utah.gov).

About the authors: This document was authored by the Utah Cannabis Research Review Board and Department of Health and Human Services staff.

About the Utah Cannabis Research Review Board: Under Utah Health Code 26B-1-420, the Cannabis Research Review Board is a board of medical research professionals and physicians who meet on a voluntary basis to review and discuss any available scientific research related to the human use of cannabis, cannabinoid product or an expanded cannabinoid product that was conducted under a study approved by an Institutional Review Board (IRB) or was conducted and approved by the federal government.

IMPORTANT NOTE: As always, in the event of significant side effects, stop use of medical cannabis until side effects have resolved, and then reduce to previous, best-tolerated dose. To avoid unwanted psychoactive side effects, “start low and go slow” especially when using cannabis products for the first time or using new dosages or types of products.

There is insufficient evidence (National Academies of Science, Engineering, and Medicine, 2017) to support or refute that short-term (8-10 week) cannabis treatment reduces inflammation or induces clinical or endoscopic response/remission in patients with active Crohn’s disease or ulcerative colitis (trials for Crohn’s disease: THC-predominant cannabis cigarettes or CBD-rich cannabis oil; trials for ulcerative colitis: CBD-predominant cannabis oral capsule or THC-predominant cannabis cigarettes)

There is limited evidence (National Academies of Science, Engineering, and Medicine, 2017) to support the conclusion that short-term (8-10 week) cannabis treatment may improve patient-reported quality-of-life in patients with active Crohn’s disease or ulcerative colitis (trials for Crohn’s disease: CBD-rich cannabis oil; trials for ulcerative colitis: CBD-predominant cannabis oral capsule or THC-predominant cannabis cigarettes).

Smoking cannabis is not permitted under the Utah health code. Any mention of smoking in this document refers to the method used for a particular study and is stated for your information only. The Department of Health and Human Services, the Cannabis Research Review Board, and the State of Utah do not promote smoking as a method of cannabis use.

The below-mentioned sections on ulcerative colitis and Crohn’s disease are adapted from the Cochrane Database Systematic Reviews.

Cannabis and cannabinoids are often promoted as treatment for many illnesses and are widely used among patients with ulcerative colitis (UC). Few studies have evaluated the use of these agents. Further, cannabis has potential for adverse

events, and the long-term consequences of cannabis and cannabinoid use in UC are unknown.

A Cochrane meta-analysis was published in Issue 6, 2019. The primary outcomes were clinical remission and relapse. Secondary outcomes included endoscopic response, quality of life, adverse events, and cannabis dependence and withdrawal effects.

Two studies met the inclusion criteria. One study compared 10 weeks of cannabidiol capsules containing up to 4.7% delta-9-tetrahydrocannabinol (THC) with placebo capsules in participants with mild to moderate UC. The starting dose of cannabidiol was 50 mg twice daily, increasing to 250 mg twice daily if tolerated. Another study compared 8 weeks of therapy with two cannabis cigarettes per day containing 0.5 g of cannabis, corresponding to 23 mg THC/day, to placebo cigarettes in participants with UC who did not respond to conventional medical treatment. The effect of cannabidiol capsules (100 mg to 500 mg daily) compared to placebo on clinical remission and response is uncertain. Clinical remission at 10 weeks was achieved by 24% of the cannabidiol group compared to 26% in the placebo group. Clinical response and Serum CRP levels were similar in both groups after 10 weeks of therapy. There may be a clinically meaningful improvement in QoL at 10 weeks. Adverse events were more frequent in cannabidiol participants compared to placebo. One hundred percent of cannabidiol participants had an adverse event, compared to 77% of placebo participants. However, these adverse events were considered to be mild or moderate in severity. Common adverse events included dizziness, disturbance in attention, headache, nausea, and fatigue. More participants in the cannabidiol group withdrew due to an adverse event than placebo participants. Withdrawals in the cannabidiol group were mostly due to dizziness. Withdrawals in the placebo group were due to worsening UC. The effect of cannabis cigarettes (23 mg THC/day) compared to placebo on mean disease activity, CRP levels, and mean fecal calprotectin levels is uncertain. After 8 weeks, the mean disease activity index score in cannabis participants was 4 compared with 8 in placebo participants. After 8 weeks, the mean change in CRP levels was similar in both groups. No serious adverse events were observed. This study did not report on clinical remission, clinical response, quality of life, adverse events, or withdrawal due to adverse events.

Conclusions. The effects of cannabis and cannabidiol on UC are uncertain. Thus, no firm conclusions regarding the efficacy and safety of cannabis or cannabidiol in adults with active UC can be drawn. There is no evidence for cannabis or cannabinoid use for

maintenance of remission in UC. Further studies with a larger number of patients are required to assess the effects of cannabis in UC patients with active and quiescent disease. Different doses of cannabis and routes of administration should be investigated. Lastly, follow-up is needed to assess the long-term safety outcomes of frequent cannabis use.

Crohn's disease (CD) is a chronic immune-mediated condition of transmural inflammation in the gastrointestinal tract, associated with significant morbidity and decreased quality of life. The endocannabinoid system provides a potential therapeutic target for cannabis, cannabinoids, and animal models have shown benefits in decreasing inflammation. However, there is also evidence to suggest transient adverse events such as weakness, dizziness, diarrhea, and an increased risk of surgery in people with CD who use cannabis.

Cochrane meta-analysis was published in Issue 6, 2019. The primary outcomes were clinical remission and relapse. Secondary outcomes included endoscopic response, quality of life, adverse events, and cannabis dependence and withdrawal effects.

Three studies met the inclusion criteria. Participants in two of the studies were adults with active Crohn's disease who had failed at least one medical treatment. One small study compared eight weeks of treatment with cannabis cigarettes containing 115 mg of delta-9-tetrahydrocannabinol (THC) to placebo cigarettes containing cannabis with the THC removed in participants with active CD. The effects of cannabis on clinical remission were unclear. A difference was observed in clinical response rates. Ninety-one percent of the cannabis group achieved a clinical response compared to 40% of the placebo group. More AEs were observed in the cannabis cigarette group compared to placebo. These AEs were considered mild in nature and included sleepiness, nausea, difficulty with concentration, memory loss, confusion, and dizziness. One small study compared cannabis oil (5% cannabidiol) to placebo oil in people with active CD. There was no difference in clinical remission rates. Forty percent of cannabis oil participants achieved remission at 8 weeks compared to 33% of the placebo participants. There was no difference in the proportion of participants who had a serious adverse event. One small study compared cannabis oil (15% cannabidiol and 4% THC) to placebo in participants with active CD. Differences in QoL scores were observed. The mean QoL score after 8 weeks of treatment was 96.3 in the cannabis oil group compared to 79.9 in the placebo group. This study did not report on clinical remission, clinical response, CRP, or AEs.

Conclusions. The effects of cannabis and cannabis oil on Crohn's disease are uncertain. Thus, no firm conclusions regarding the efficacy and safety of cannabis and cannabis oil in adults with active Crohn's disease can be drawn. The effects of cannabis or cannabis oil in quiescent Crohn's disease have not been investigated. Further studies with larger numbers of participants are required to assess the potential benefits and harms of cannabis in Crohn's disease. Future studies should assess the effects of cannabis in people with active and quiescent Crohn's disease. Different doses of cannabis and delivery modalities should be investigated.

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DISCLAIMER

The following information on the use of medical cannabis serves as a suggested use guide for those participating in the Utah Medical Cannabis Program. This document has been vetted and approved by the Utah Cannabis Research Review Board under Utah Health Code 26B-1-420.

This document is a summary of available peer-reviewed literature concerning potential therapeutic uses and harmful effects of cannabis and cannabinoids. With the ongoing nature of cannabis and cannabinoid research, it is not meant to be complete or comprehensive and should be used as a limited complement to other reliable sources of information. This document is not a systematic review or meta-analysis of the literature and has not rigorously evaluated the quality and weight of the available evidence. There is a lack of controlled clinical trials yielding high level evidence of predictable therapeutic benefit for any given condition other than those for FDA approved formulations. This document includes warnings and risks related to the use of cannabis including cannabis use disorder, potentially irreversible brain damage/mental illness, and legal liability for DUI and potential for adverse work-related consequences.

All patrons participating in the Utah Medical Cannabis Program are advised to use this document and any such document produced from this original document as informational and educational. The use of medical cannabis is at one's own risk. **Medical cannabis is NOT a first line therapy for most medical conditions.**

The information in this document is intended to help as far as available data allows Utah health care decision-makers, health care professionals, health systems leaders, and Utah Medical Cannabis patients to make well-informed decisions and thereby improve the quality of health care outcomes in patients using medical cannabis use. While patients and others may access this document, the document is made available for informational purposes only and no representations or warranties are made with respect to its fitness for any particular purpose. The information in this document should not be used as a substitute for professional medical advice or as a substitute for the application of clinical judgment in respect of the care of a particular patient or other professional judgment in any decision-making process.

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